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APPLICATION NO.	FILING DATE	FIRST NAMED INVENTOR	ATTORNEY DOCKET NO.	CONFIRMATION NO.	
09/942,586	08/31/2001	James Hager	571-737	9428	
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H. Samuel Frost			EXAM	EXAMINER	
Bereskin & Parr Box 401			JOHNSTON,	JOHNSTON, PHILLIP A	
40 King Street West Toronto, ON M5H 3Y2			ART UNIT	PAPER NUMBER	
CANADA			2881		
			DATE MAILED: 01/02/2003	DATE MAILED: 01/02/2003	

Please find below and/or attached an Office communication concerning this application or proceeding.

			<u> </u>			
•	_	Application No.	Applicant(s)			
. Office Action Summary		09/942,586	HAGER, JAMES			
	Office Action Summary	Examiner	Art Unit			
	The MAILING DATE of this communication com	Phillip A Johnston	2881			
Period fo	The MAILING DATE of this communication appears on the cover sheet with the correspondence address Period for Reply					
A SHORTENED STATUTORY PERIOD FOR REPLY IS SET TO EXPIRE 3 MONTH(S) FROM THE MAILING DATE OF THIS COMMUNICATION. - Extensions of time may be available under the provisions of 37 CFR 1.136(a). In no event, however, may a reply be timely filed after SIX (6) MONTHS from the mailing date of this communication. - If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely. - If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication. - Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). - Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b).						
1)	Responsive to communication(s) filed on					
2a)□	• • • • • • • • • • • • • • • • • • • •	— · s action is non-final.				
3)□	Since this application is in condition for allowar		osecution as to the merits is			
closed in accordance with the practice under <i>Ex parte Quayle</i> , 1935 C.D. 11, 453 O.G. 213. Disposition of Claims						
4)⊠ Claim(s) <u>1-23</u> is/are pending in the application.						
4a) Of the above claim(s) is/are withdrawn from consideration.						
5)	Claim(s) is/are allowed.					
6)⊠ Claim(s) <u>1-23</u> is/are rejected.						
7)	7) Claim(s) is/are objected to.					
8) Claim(s) are subject to restriction and/or election requirement.						
Application Papers						
9) The specification is objected to by the Examiner.						
10)⊠ T	10) \boxtimes The drawing(s) filed on <u>31 August 2001</u> is/are: a) \boxtimes accepted or b) \square objected to by the Examiner.					
Applicant may not request that any objection to the drawing(s) be held in abeyance. See 37 CFR 1.85(a).						
11) The proposed drawing correction filed on is: a) approved b) disapproved by the Examiner.						
If approved, corrected drawings are required in reply to this Office action.						
12) The oath or declaration is objected to by the Examiner.						
Priority under 35 U.S.C. §§ 119 and 120						
13) Acknowledgment is made of a claim for foreign priority under 35 U.S.C. § 119(a)-(d) or (f).						
a) All b) Some * c) None of:						
1. Certified copies of the priority documents have been received. 2. Certified copies of the priority documents have been received in Application No.						
2. Certified copies of the priority documents have been received in Application No						
 3. Copies of the certified copies of the priority documents have been received in this National Stage application from the International Bureau (PCT Rule 17.2(a)). * See the attached detailed Office action for a list of the certified copies not received. 						
14)⊠ Acknowledgment is made of a claim for domestic priority under 35 U.S.C. § 119(e) (to a provisional application).						
a) The translation of the foreign language provisional application has been received. 15) Acknowledgment is made of a claim for domestic priority under 35 U.S.C. §§ 120 and/or 121.						
Attachment(s)						
2) Notice	of References Cited (PTO-892) of Draftsperson's Patent Drawing Review (PTO-948) ation Disclosure Statement(s) (PTO-1449) Paper No(s)	5) Notice of Informal Pa	PTO-413) Paper No(s) atent Application (PTO-152)			

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Detailed Action

Claims Rejection - 35 U.S. C. 102

- 1. The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:
 - (b) the invention was patented or described in a printed publication in this or a foreign country or in public use or on sale in this country, more than one year prior to the date of application for patent in the United States.
- 2. Claims 1-5, and 7-10 are rejected under 35 U.S.C. 102(b) as being clearly anticipated by Whitehouse, U. S. Patent No. 6,188, 066.

Whitehouse discloses a multipole ion guide, which is used to transport ions entering vacuum from an API source to non-dispersion type mass analyzers. A range of ion mass to charge (m/z) can be efficiently transmitted through a multipole ion guide provided the ion guide operating stability region is set to pass those values of m/z. If an ion with a given mass to charge ratio falls within the operating stability region set for a multipole ion guide, the ion will be effectively trapped from drifting to far in the off axis direction but is free to move in the direction of ion guide axis. See Column 3, line 51-61. The offset potential of the multipole lens, that is the DC voltage applied uniformly to all the rods on

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which the AC and alternate polarity DC rod potentials are floated and referenced is one variable that can to be used to set the energy of ions transmitted through the multipole ion guide. See Column 4, line 10-14.

Whitehouse teaches that a valuable feature of multipole ion guides when operated in higher background pressures is that ions traversing the length of the ion guide experience a number of collisions with the background gas resulting in the cooling of the ion kinetic energy, as recited in Claim 3. As the ions enter the multipole ion guide and are transmitted through it, the RF or combined RF-DC field effectively traps the ions from dispersing in the radial direction due to collisions with the background gas yet permits movement of ions in the axial direction often driven by the gas dynamics. Ions which experience a number of low energy collisions with the neutral gas within the multipole rod assembly have their kinetic energy reduced resulting in a narrowing of the ion energy spread for those ions which exit the multipole rods, as recited in Claim 2. The number of collisions an ion experiences as it travels the length of the ion guide is a function of the rod length and the background pressure inside the rods. If the relative voltages of the capillary exit lens 50, ring lens 51, skimmer 47 and the multipole ion guide 40 DC offset potential remain the same then the ions entering the multipole ion guide will have similar energy spreads and will be transmitted to the exit of the multipole ion guide with the same efficiency. See column 13, line 9-30.

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Whitehouse further shows in FIG. 11, a three vacuum pumping stage API/ion trap system, as recited in Claim 5, where a multipole ion guide begins in vacuum pumping stage 148 and extends continuously through the three vacuum pumping stages. See Column 19, line 24-28. In addition, the multipole ion guide 165 can efficiently transport ions through gradient in background gas pressure. As was shown in FIGS. 8a, b and c, the ion energy spread is reduced due to ion collisional cooling with the background gas. Higher trapping efficiency can be achieved with ions entering ion trap 154 when the ions have a narrow energy spread. Increased trapping efficiencies result in higher signal sensitivity for a given ion current entering trap 154. The ability to selectivity cutoff a range of m/z transmission through multipole ion guide 165 can also be used to increase sensitivity in ion trap mass spectrometers. Ion traps must first trap ions and then conduct a mass analysis on a packet of trapped ions. The trap can only hold a limited number of ions before it suffers from space charge effects, which can shift measured m/z values and deteriorate resolution. See Column 20, line 19-33.

3. Claims 6, 11-14, and 22 are rejected under 35 U.S.C. 102(b) as being clearly anticipated by Whitehouse, U. S. Patent No. 6,188, 066, as applied above.

Whitehouse teaches that the ability of a multipole ion guide to <u>deliver an</u> ion beam with low energy spread and where the mean energy and m/z range can be adjusted into a mass analyzer can be used to improve the

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performance of an API/Time-Of-Flight, <u>API/Ion Trap</u> and API/FT-ICR mass spectrometer systems, as recited in Claims 6, 8,10-14, and 22. See Column 4, line 16-21.

Whitehouse further discloses an Electrospray ion source interfaced to a quadrupole mass spectrometer is diagramed in FIG. 1. The system shown includes four vacuum pumping stages and a static voltage lens configuration to transfer ions through the first three pumping stages and focus them into the entrance of quadrupole mass spectrometer 18. An electrostatic field is applied between the capillary exit 8, the ring lens 9 and the first skimmer 10 to electrostatically focus and accelerate ions through the skimmer 10 orifice 11 and on into the second vacuum stage 12.

Electrostatic potentials are maintained between skimmers 10 and 13 and a portion of the ions passing through skimmer 10 are focused through orifice 22 in skimmer 13 into the third vacuum pumping stage 20. Potentials are set on electrostatic lens elements 14, 15 and 16 to focus ions through aperture 17 after which they pass into the quadrupole mass filter 18 located in the fourth pumping stage 24. See Column 7, line 57-63, and Column 8, line 18-33.

4. Claims 15-21, and 23 are also rejected under 35 U.S.C. 102(b) as being clearly anticipated by Whitehouse, U. S. Patent No. 6,188, 066, as applied above.

Whitehouse discloses that the ion energy relative to the quadrupole mass filter offset voltage is established by a combination of acceleration energy imparted by the expanding carrier gas and the electrostatic potentials applied. The capillary exit 8

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potential relative to the ring electrode 9 and skimmer electrode 10 can be set high enough to cause Collisional Induced Dissociation (CID) which can affect the energy and energy spread of the parent and fragment ions. See Column 8, line 37-44.

Whitehouse also teaches that the collisional cross sections of ions generated by API sources are not always known, however. In particular, the cross section of <u>multiply charged ions</u> which can be produced in the electrospray ion source are not always known. Consequently, for a given multipole ion guide configuration, values of a_n and q_n where efficient ion transmission through the multipole lens can be achieved must be mapped for any given m/z and background pressure combination encountered. See Column 10, line 33-42.

Whitehouse further discloses that the ability of the multipole ion guide to selectively transmit a range of m/z values while cutting off the transmission of m/z outside that range can be used to increase the duty cycle and detector sensitivity in an API/TOF system. If a DC potential is applied to the poles where each adjacent pole has opposite DC polarity, the a_n and q_n values can be selected so that the multipole ion guide will pass a narrower range of m/z. See Column 17, line 24-36.

Whitehouse further teaches that one variable which effects duty cycle in a TOF instrument is the repetition rate at which ions are pulsed in the flight tube, accelerated and detected. Assuming that the pulsing region 135 can refill between pulses, that is the primary ion beam 134 energy is set to satisfy this criteria, the pulsing repetition rate will be limited by the fastest flight time of the lowest m/z ion and

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through flight tube 127 to the detector. Any overlap of ions from one pulsed packet to the next will increase the difficulty in interpreting the resulting mass spectrum. If ions of lower or higher m/z were not of interest in a given analysis, those ions could be preventing from entering the pulsing region by selecting an appropriate a_n and q_n value for the multipole rod 18 operation. By reducing the arrival time spread of an ion packet as it travels down the flight tube, the time in between pulses can be reduced resulting in an increase in duty cycle. See Column 17, line 46-62.

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Conclusion

5. Any inquiry concerning this communication or earlier communications should be directed to Phillip Johnston whose telephone number is (703) 305-7022. The examiner can normally be reached on Monday-Friday from 8:00 am to 5:30 pm.

If attempts to reach the examiner by telephone are unsuccessful, the examiners supervisor John Lee can be reached at (703) 308-4116. The $\frac{1}{15}$ $\frac{703}{15}$ $\frac{8}{15}$ $\frac{2}{15}$ $\frac{2}{15}$ $\frac{2}{15}$ fax phone numbers are (703) 308-2864 and (703) 308-7721.

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